Amendment to the Claims:

This listing of claims will replace all previous versions, and listings, of claims in this application.

Listing of Claims:

1. (currently amended) A compound having the formula I of the following formula

$$\begin{array}{c|c}
Z & NH_2 \\
\hline
 & X & Q \\
\hline
 & (R^4)_n
\end{array}$$

$$\begin{array}{c} (H) \\ Z \\ NH_2 \\ Y \\ Q \\ (R^4)_n \end{array}$$

wherein:

Z is N;

Y is CONR⁵, NR⁵CO, SO₂NR⁵, NR⁵SO₂, CH₂NR⁵, NR⁵CONR⁵, CH₂CO, CO or CH₂O;

X is N;

P is phenyl;

Q is C_1 -6alkyl, C_2 -6alkenyl or C_2 -6alkynyl;

R is C_{0-6} alkyl(SO_2) NR^1R^2 ;

 R^4 -and R^2 -are independently selected from hydrogen, C_4 - $_6$ alkyl, C_2 - $_6$ alkenyl, C_2 - $_6$ alkynyl, C_6 - $_6$ alkyl C_3 - $_6$ cycloalkyl, C_6 - $_6$ alkylheterocycloalkyl, C_4 - $_6$ alkyl C_3 - $_6$ cycloalkyl, C_6 - $_6$ alkylheteroaryl, wherein any C_4 - $_6$ alkyl, C_2 - $_6$ alkenyl, C_2 - $_6$ alkynyl, C_6 - $_6$ alkylheterocycloalkyl, C_6 - $_6$ alkylaryl, C_6 - $_6$ alkylheteroaryl may be substituted by one or more C_6 :

R¹ and R² [[may]] together form a substituted 5 or 6 membered heterocyclic ring containing one or more heteroatoms independently selected from N, O or S, which heterocyclic ring may be optionally substituted by A;

 $R^{2} \text{ is independently selected from halogen, nitro, CHO, $C_{0.6}$ alkylCN, $OC_{1.6}$ alkylCN, $C_{0.6}$ alkylOR$^{6}, $OC_{1.6}$ alkylOR$^{6}, fluoromethyl, difluoromethyl, trifluoromethyl, fluoromethoxy, difluoromethoxy, trifluoromethoxy, $C_{0.6}$ alkylNR^{6}R^{7}, $OC_{1.6}$ alkylNR^{6}R^{7}, $OC_{1.6}$ alkylNR^{6}R^{7}, NR^{6}OR$^{7}, $C_{0.6}$ alkylCO_{2}R$^{6}, $C_{0.6}$ alkylCO_{2}R$^{6}, $C_{0.6}$ alkylCONR^{6}R^{7}, $OC_{1.6}$ alkylOC_{1.6}$ alkylNR$^{6}(CO)R$^{7}, $OC_{1.6}$ alkylCONR^{6}R^{7}, $OC_{1.6}$ alkylNR$^{6}(CO)R$^{7}, $O(CO)NR$^{6}R$^{7}, NR^{6}(CO)NR^{6}R^{7}, $O(CO)NR$^{6}R$^{7}, $O(CO)R$^{6}, $O(CO)R$^{6}, $C_{0.6}$ alkylCOR$^{6}, $OC_{1.6}$ alkylCOR$^{6}, NR^{6}(CO)(CO)NR^{6}R^{7}, SR^{6}, $C_{0.6}$ alkyl(SO_{2})NR^{6}R^{7}, $OC_{1.6}$ alkylNR$^{6}(SO_{2})R$^{7}, $OC_{0.6}$ alkylNR$^{6}(SO_{2})NR$^{6}R$^{7}, $OC_{0.6}$ alkylNR$^{6}(SO_{2})NR$^{6}R$^{7}, $OC_{0.6}$ alkylNR$^{6}(SO_{2})NR$^{6}R$^{7}, $OC_{0.6}$ alkylNR$^{6}(SO_{2})NR$^{6}R$^{7}, $OC_{0.6}$ alkylNR$^{6}(SO_{2})R$^{7}, $OC_{0.6}$ alkylNR$^{7}, $OC_{0.6}$ al$

R⁴ is independently selected from halogen, nitro, CHO, CN, OC₁₋₆alkylCN, OR⁶, OC₁₋₆alkylOR⁶; fluoromethyl, difluoromethyl, trifluoromethyl, fluoromethoxy, difluoromethoxy, trifluoromethoxy, NR⁶R⁷, OC₁₋₆alkylNR⁶R⁷, NR⁶OR⁷, CO₂R⁶, OC₁₋₆alkylCO₂R⁶, CONR⁶R⁷; OC₁₋₆alkylNR⁶(CO)R⁷, NR⁶(CO)R⁷, O(CO)NR⁶R⁷, NR⁶(CO)OR⁷; NR⁶(CO)OR⁶, O(CO)OR⁶, O(CO)R⁶, OC₁₋₆alkylCOR⁶, NR⁶(CO)(CO)R⁶; NR⁶(CO)(CO)NR⁶R⁷, SR⁶, (SO₂)NR⁶R⁷, OC₁₋₆alkylNR⁶(SO₂)R⁷, OC₀₋₆alkyl(SO₂)NR⁶R⁷, (SO)NR⁶R⁷, OC₁₋₆alkylSO₂R⁶, NR⁶(SO₂)NR⁶R⁷, NR⁶(SO)R⁷, OC₁₋₆alkylNR⁶(SO)R⁷, OC₀₋₆alkylSO₂R⁶, SO₂R⁶, SO₂R⁶, SOR⁶, C₂₋₆cycloalkyl, CN, OR⁶, CONR⁶R⁷, NR⁶COR⁷, (SO)NR⁶R⁷, SO₂R⁶, phenyl, a 5 or 6 membered heteroaromatic ring containing one or more heteroatoms independently selected from N, O, or S, or a 5 or 6 membered heterocyclic ring containing one or more heteroatoms independently selected from N, O, or S which heterocyclic group may be saturated or unsaturated, and said phenyl ring or 5 or 6 membered heteroaromatic ring or 5 or 6 membered heteroaromatic ring or 5 or 6 membered heterocyclic ring may optionally be fused with a 5 or 6 membered saturated, partially saturated or unsaturated ring containing atoms independently

selected from C, N, O or S wherein any C₃₋₆eycloalkyl, phenyl, 5 or 6 membered heteroaromatic ring with one or two heteroatoms selected independently from N, O, or S or a 5 or 6 membered heterocyclic containing one or two heteroatoms selected independently from N, O, or S; may be optionally be substituted by one or more A;

m is 0:

n is [[0,]] 1, 2, 3 or 4;

R⁵ is hydrogen or C₁-6alkyl;

 R^6 and R^7 are independently selected from hydrogen, <u>or</u> C_{1^-6} alkyl, C_{2^-6}

 R^6 -and R^7 -may together form a substituted 5 or 6 membered heterocyclic ring containing one or more heteroatoms independently selected from N, O or S, which heterocyclic ring may be optionally substituted by A and wherein a CH_2 group may optionally be replaced by a CO group; R^8 -and R^9 -are independently selected from hydrogen, C_1 -alkyl, C_2 -alkenyl, C_2 -alkynyl, C_0 -alkyl C_3 -acycloalkyl, C_0 -alkylaryl and C_0 -alkylheteroaryl;

R⁸ and R⁹ may together form a 5 or 6 membered heterocyclic ring containing one or more heteroatoms selected from N, O or S, which heterocyclic ring may be optionally substituted by A;

$$\begin{split} &R^{1\theta}\text{-is hydrogen, }C_{1}\text{-}_{6}\text{alkyl, }C_{2}\text{-}_{6}\text{alkenyl, }C_{2}\text{-}_{6}\text{alkynyl, }C_{\theta}\text{-}_{6}\text{alkyl}C_{3}\text{-}_{6}\text{cycloalkyl, }\\ &C_{\theta}\text{-}_{6}\text{alkylaryl, }C_{\theta}\text{-}_{6}\text{alkylheteroaryl or }C_{1}\text{-}_{6}\text{alkylNR}^{8}R^{9}\text{;} \end{split}$$

 R^{11} is C_{1-6} alky $1NR^{8}R^{9}$;

R¹⁰-and R¹¹-may together form a 5 or 6 membered heterocyclic ring containing one or more heteroatoms selected from N, O or S, which heterocyclic ring may be optionally substituted by A;

 R^{12} is a 5 or 6 membered heterocyclic ring containing one or more heteroatoms independently selected from N, O or S, which heterocyclic ring may be optionally substituted by Λ ; wherein any C_1 -6alkyl, C_2 -6alkenyl, C_2 -6alkynyl, C_0 -6alkyl C_3 -6cycloalkyl, C_0 -6alkylheterocycloalkyl, C_0 -6alkylheterocycloalkyl, C

Application No. 10/539,545 Response to Office action dated 09/22/2008

 $\frac{CONR^6R^7,NR^6(CO)R^6,O(CO)R^6,COR^6,SR^6,(SO_2)NR^6R^7,(SO)NR^6R^7,SO_2R^6,SO_2R^6-orSOR^6;}{SOR^6};$

as a free base or a pharmaceutically acceptable salt, solvate or solvate of a salt thereof.

Claim 2 (cancelled).

- 3. (previously presented) A compound according to claim 1, wherein R^1 and R^2 in C_{0-6} alkyl(SO_2)N R^1R^2 together form a substituted 5 or 6 membered heterocyclic ring containing one or more heteroatoms selected from N, O or S.
- 4. (original) A compound according to claim 3, wherein said heterocyclic ring comprises one or more N heteroatoms and said heterocyclic ring is optionally substituted by A, preferably a C_{1-6} alkyl.
- 5. (previously presented) A compound according to any one of claims 1, 3 or 4, wherein Y is CONR⁵; R⁵ is hydrogen; Q is C₁₋₆alkyl; R⁴ is selected from: phenyl, 5 or 6 membered heteroaromatic ring containing one or more heteroatoms independently selected from N, O, or S or a 5 or 6 membered heterocyclic ring containing one or two heteroatoms selected independently from N, O, or S which heterocyclic group may be saturated or unsaturated, CN, OR⁶, SO₂R⁶, NR⁶(CO)R⁷, (SO₂)NR⁶R⁷, and CONR⁶R⁷; and n is 1; said phenyl or 5 or 6 membered heterocyclic ring optionally substituted by A.
- 6. (original) A compound according to claim 5, wherein A is selected from OR^6 , C_{1-6} alkyl, oxo (=O) and nitro; and R^6 and/or R^7 is selected from C_{1-6} alkyl and hydrogen.

- 7. (currently amended) A compound which is
- 3-Amino-*N*-(2-cyanoethyl)-6-[4-(pyrrolidin-1-ylsulfonyl)phenyl]pyrazine-2-carboxamide;
- 3-Amino-*N*-(3-amino-3-oxopropyl)-6-[4-(pyrrolidin-1-ylsulfonyl)phenyl]pyrazine-2-carboxamide;
- 3-Amino-N-(2-nitrobenzyl)-6-[4-(pyrrolidin-1-ylsulfonyl)phenyl]pyrazine-2-carboxamide;
- 3-Amino-*N*-(2-methoxybenzyl)-6-[4-(pyrrolidin-1-ylsulfonyl)phenyl]pyrazine-2-carboxamide;
- 3-Amino-*N*-(3-morpholin-4-ylpropyl)-6-[4-(pyrrolidin-1-ylsulfonyl)phenyl]pyrazine-2-carboxamide;
- 3-Amino-*N*-[3-(4-methylpiperazin-1-yl)propyl]-6-[4-(pyrrolidin-1-ylsulfonyl)phenyl]pyrazine-2-carboxamide;
- as a free base or a pharmaceutically acceptable salt, solvate or solvate of a salt thereof;
- 3-Amino-*N*-(2-morpholin-4-ylethyl)-6-[4-(pyrrolidin-1-ylsulfonyl)phenyl]pyrazine-2-carboxamide hydrochloride;
- 3-Amino-*N*-[2-(1*H*-imidazol-4-yl)ethyl]-6-[4-(pyrrolidin-1-ylsulfonyl)phenyl]pyrazine-2-carboxamide hydrochloride;
- 3-Amino-*N*-[3-(1*H*-imidazol-1-yl)propyl]-6-[4-(pyrrolidin-1-ylsulfonyl)phenyl]pyrazine-2-carboxamide hydrochloride;
- 3-Amino-6-{4-[(4-methylpiperazin-1-yl)sulfonyl]phenyl}-*N*-(2-thien-2-ylethyl)pyrazine-2-carboxamide hydrochloride;
- 3-Amino-6-{4-[(4-methylpiperazin-1-yl)sulfonyl]phenyl}-*N*-(thien-2-ylmethyl)pyrazine-2-carboxamide hydrochloride;
- 3-Amino-*N*-(2-methoxyethyl)-6-{4-[(4-methylpiperazin-1-yl)sulfonyl]phenyl}pyrazine-2-carboxamide hydrochloride;
- 3-Amino-*N*-(3-methoxypropyl)-6-{4-[(4-methylpiperazin-1-yl)sulfonyl]phenyl}pyrazine-2-carboxamide hydrochloride;
- 3-Amino-6-{4-[(4-methylpiperazin-1-yl)sulfonyl]phenyl}-*N*-[3-(2-oxopyrrolidin-1-yl)propyl]pyrazine-2-carboxamide hydrochloride;
- 3-Amino-*N*-(cyanomethyl)-6-{4-[(4-methylpiperazin-1-yl)sulfonyl]phenyl}pyrazine-2-carboxamide dihydrochloride;

- 3-Amino-6-[4-[(4-methyl-1-piperazinyl)sulfonyl]phenyl]-*N*-[2-(1*H*-pyrrol-1-yl)ethyl]-2-pyrazinecarboxamide hydrochloride;
- 3-Amino-6-[4-[(4-methyl-1-piperazinyl)sulfonyl]phenyl]-*N*-[2-(methylsulfonyl)ethyl]-2-pyrazinecarboxamide hydrochloride;
- *N*-[2-(Acetylamino)ethyl]-3-amino-6-[4-[(4-methyl-1-piperazinyl)sulfonyl]phenyl]-2-pyrazinecarboxamide hydrochloride;
- 3-Amino-6-[4-[(4-methyl-1-piperazinyl)sulfonyl]phenyl]-*N*-[2-(2-oxo-1-imidazolidinyl)ethyl]-2-pyrazinecarboxamide hydrochloride;
- 3-Amino-*N*-[2-(aminosulfonyl)ethyl]-6-[4-[(4-methyl-1-piperazinyl)sulfonyl]phenyl]-2-pyrazinecarboxamide hydrochloride;

or as a free base or an alternative pharmaceutically acceptable salt, solvate or solvate of a salt thereof.

8. (previously presented) A pharmaceutical formulation comprising as active ingredient a therapeutically effective amount of the compound according to any one of claims 1 or 7 in association with pharmaceutically acceptable carriers or diluents.

Claims 9 to 16. (Cancelled)

17. (withdrawn) A method of prevention and/or treatment of conditions associated with glycogen synthase kinase-3, comprising administering to a mammal, including man in need of such prevention and/or treatment, a therapeutically effective amount of a compound of formula **I** as defined in any one of claims 1 or 7.

18. (withdrawn) A method of prevention and/or treatment of dementia, Alzheimer's Disease, Parkinson's Disease, Frontotemporal dementia Parkinson's Type, Parkinson dementia complex

of Guam, HIV dementia, diseases with associated neurofibrillar tangle pathologies and dementia pugilistica, comprising administering to a mammal, including man in need of such prevention and/or treatment, a therapeutically effective amount of a compound of formula **I** as defined in any one of claims 1 or 7.

- 19. (withdrawn) The method according to claim 18, wherein the prevention and/or treatment is for Alzheimer's Disease.
- 20. (withdrawn) A method of prevention and/or treatment of amyotrophic lateral sclerosis, corticobasal degeneration, Down syndrome, Huntington's Disease, postencephalitic parkinsonism, progressive supranuclear palsy, Pick's Disease, Niemann-Pick's Disease, stroke, head trauma and other chronic neurodegenerative diseases, Bipolar Disease, affective disorders, depression, schizophrenia, cognitive disorders, hair loss and contraceptive medication, Type I and Type II diabetes, diabetic neuropathy and diabetes related disorders, comprising administering to a mammal, including man in need of such prevention and/or treatment, a therapeutically effective amount of a compound of formula I as defined in any one of claims 1 or 7.
- 21. (withdrawn) The method according to claim 18, wherein the prevention and/or treatment is of Type I or Type II diabetes, diabetic neuropathy or diabetes related disorders.
- 22. (withdrawn) A method of prevention and/or treatment of predemented states, Mild Cognitive Impairment, Age-Associated Memory Impairment, Age-Related Cognitive Decline, Cognitive Impairment No Dementia, mild cognitive decline, mild neurocognitive decline, Late-Life Forgetfulness, memory impairment and cognitive impairment, vascular dementia, dementia with Lewy bodies, Frontotemporal dementia and androgenetic alopecia, comprising administering to a

mammal, including man in need of such prevention and/or treatment, a therapeutically effective amount of a compound of formula I as defined in any one of claims 1 or 7.

23. (withdrawn) A process for the preparation of a compound of formula **I** according to claim 1, wherein Y, X, Z, P, Q, R, R¹, R², R³, R⁴, R⁵, R⁶, R⁷, R⁸, R⁹, R¹⁰, R¹¹, R¹², A, m and n are defined as in formula **I**, comprising of de-halogen coupling of a compound of formula **IV** with an appropriate aryl species;

to give a compound of formula I.

24. (withdrawn) A process for the preparation of a compound of formula I according to claim 1, wherein Y, X, Z, P, Q, R, R¹, R², R³, R⁴, R⁵, R⁶, R⁷, R⁸, R⁹, R¹⁰, R¹¹, R¹², A, m and n are defined as in formula I, comprising reacting of a compound of formula XXII:

Polystyrene
$$R \xrightarrow{P} O OH$$

$$(XXII)$$

$$R \xrightarrow{P} (R^3)_m$$

$$(XXII)$$

$$(I)$$

wherein the reaction is being performed by activation of a compound of formula **XXII** by treatment with a coupling agent or with an acyl halide reagent followed by treatment with the appropriate amine, followed by cleavage of the solid phase moiety by treatment with an suitable acid in a suitable solvent, and where the reaction temperature is between 0 °C and reflux, to give a compound of formula **I**.

25 and 26. (cancelled)

27. (withdrawn) A compound which is

4-(Pyrrolidin-1-ylsulfonyl)phenylboronic acid;

4-[(4-Methylpiperazin-1-yl)sulfonyl]phenylboronic acid;

as a free base or a salt, solvate or solvate of a salt thereof.

28. (withdrawn) A compound of formula IV

Hal
$$X \longrightarrow X^{NH_2}$$

$$Q \longrightarrow (R^4)_r$$

$$(IV)$$

wherein

Y is CONR⁵, NR⁵CO, SO₂NR⁵, NR⁵SO₂, CH₂NR⁵ NR⁵CONR⁵, CH₂CO, CO or CH₂O;

X is CH or N;

Z is N;

Q is C_1 -6alkyl, C_2 -6alkenyl or C_2 -6alkynyl;

R⁴ is independently selected from halogen, nitro, CHO, CN, OC₁₋₆alkylCN, OR⁶, OC₁₋₆alkylOR⁶, fluoromethyl, difluoromethyl, trifluoromethyl, fluoromethoxy, difluoromethoxy, trifluoromethoxy, NR⁶R⁷, OC₁₋₆alkylNR⁶R⁷, NR⁶OR⁷, CO₂R⁶, OC₁₋₆alkylCO₂R⁶, CONR⁶R⁷,

OC₁₋₆alkylCONR⁶R⁷, OC₁₋₆alkylNR⁶(CO)R⁷, NR⁶(CO)R⁷, O(CO)NR⁶R⁷, NR⁶(CO)OR⁷, NR⁶(CO)OR⁶, O(CO)R⁶, O(CO)R⁶, OC₁₋₆alkylCOR⁶, NR⁶(CO)(CO)R⁶, NR⁶(CO)(CO)NR⁶R⁷, SR⁶, (SO₂)NR⁶R⁷, OC₁₋₆alkylNR⁶(SO₂)R⁷, OC₀₋₆alkyl(SO₂)NR⁶R⁷, (SO)NR⁶R⁷, OC₁₋₆alkyl(SO)NR⁶R⁷, SO₃R⁶, NR⁶(SO₂)NR⁶R⁷, NR⁶(SO)R⁷, OC₁₋₆alkylNR⁶(SO)R⁷, OC₀₋₆alkylSO₂R⁶, SO₂R⁶, SO₂R⁶, SOR⁶, C₃₋₆cycloalkyl, phenyl, a 5 or 6 membered heteroaromatic ring containing one or more heteroatoms independently selected from N, O, or S, or a 5 or 6 membered heterocyclic ring containing one or more heteroatoms independently selected from N, O, or S which heterocyclic group may be saturated or unsaturated, and said phenyl ring or 5 or 6 membered heteroaromatic ring or 5 or 6 membered heterocyclic ring may optionally be fused with a 5 or 6 membered saturated, partially saturated or unsaturated ring containing atoms independently selected from C, N, O or S wherein any C₃₋₆cycloalkyl, phenyl, 5 or 6 membered heteroaromatic ring with one or two heteroatoms selected independently from N, O, or S or a 5 or 6 membered heterocyclic ring containing one or two heteroatoms selected independently from N, O, or S; may be optionally be substituted by one or more A;

R⁵ is hydrogen or C₁-6alkyl

 R^6 and R^7 are independently selected from hydrogen, $C_{1\text{-}6}$ alkyl, $C_{2\text{-}6}$ alkenyl, $C_{2\text{-}6}$ alkynyl, $C_{0\text{-}6}$ alkyl $C_{3\text{-}6}$ cycloalkyl, $C_{0\text{-}6}$ alkylaryl, $C_{0\text{-}6}$ alkylheteroaryl and $C_{1\text{-}6}$ alkyl NR^8R^9 ;

 R^6 and R^7 may together form a substituted 5 or 6 membered heterocyclic ring containing one or more heteroatoms independently selected from N, O or S, which heterocyclic ring may be optionally substituted by A and wherein a CH_2 group may optionally be replaced by a CO group; R^8 and R^9 are independently selected from hydrogen, $C_{1\text{-}6}$ alkyl, $C_{2\text{-}6}$ alkenyl, $C_{2\text{-}6}$ alkynyl, $C_{0\text{-}6}$ alkyl $C_{3\text{-}6}$ cycloalkyl, $C_{0\text{-}6}$ alkylaryl and $C_{0\text{-}6}$ alkylheteroaryl;

R⁸ and R⁹ may together form a 5 or 6 membered heterocyclic ring containing one or more heteroatoms selected from N, O or S, which heterocyclic ring may be optionally substituted by A;

Hal is halogen;

n is 0, 1, 2, 3 or 4;

A is halogen, oxo (=O), nitro, CHO, CN, OR⁶, C₁₋₆alkyl, C₂₋₆alkenyl, C₂₋₆alkynyl, C₀₋₆alkylC₃₋₆cycloalkyl, fluoromethyl, difluoromethyl, trifluoromethyl, fluoromethoxy,

difluoromethoxy, trifluoromethoxy, C_{0-6} alkylNR⁶R⁷, OC_{1-6} alkylNR⁶R⁷, CO_2 R⁸, $CONR^6$ R⁷, NR⁶(CO)R⁶, O(CO)R⁶, COR⁶, SR⁶, (SO₂)NR⁶R⁷, (SO)NR⁶R⁷, SO₃R⁶, SO₂R⁶ or SOR⁶; as a free base or a salt, solvate or solvate of a salt thereof.

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29. (withdrawn) A compound according to claim 28, wherein Y is CONR<sup>5</sup>;
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X is N:

Q is C_{1-6} alkyl;

R⁴ is independently selected from CN, OR⁶, a 5 or 6 membered heteroaromatic ring containing one or more heteroatoms independently selected from N, O, or S, or a 5 or 6 membered heterocyclic ring containing one or more heteroatoms independently selected from N, O, or S which heterocyclic group may be saturated or unsaturated, wherein any 5 or 6 membered heterocyclic ring containing one or two heteroatoms selected independently from N, O, or S; may be optionally be substituted by A;

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R<sup>5</sup> is hydrogen;
R<sup>6</sup> is, C<sub>1</sub>-<sub>6</sub>alkyl;
n is 1;
A is oxo (=O);
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as a free base or a salt, solvate or solvate of a salt thereof.

- 30. (withdrawn) A compound which is
- 3-Amino-6-bromo-*N*-(2-morpholin-4-ylethyl)pyrazine-2-carboxamide;
- 3-Amino-6-bromo-*N*-[2-(1*H*-imidazol-4-yl)ethyl]pyrazine-2-carboxamide;
- 3-Amino-6-bromo-*N*-[3-(1*H*-imidazol-1-yl)propyl]pyrazine-2-carboxamide;
- 3-Amino-6-bromo-*N*-(2-thien-2-ylethyl)pyrazine-2-carboxamide;
- 3-Amino-6-bromo-*N*-(thien-2-ylmethyl)pyrazine-2-carboxamide;
- 3-Amino-6-bromo-*N*-(2-methoxyethyl)pyrazine-2-carboxamide;
- 3-Amino-6-bromo-*N*-(3-methoxypropyl)pyrazine-2-carboxamide;

3-Amino-6-bromo-*N*-[3-(2-oxopyrrolidin-1-yl)propyl]pyrazine-2-carboxamide;

3-Amino-6-bromo-*N*-(cyanomethyl)pyrazine-2-carboxamide;

as a free base or a salt, solvate or solvate of a salt thereof.

31. (withdrawn) A compound of formula XXII

$$\begin{array}{c} & & & \\ & &$$

(XXII)

wherein:

Z is N;

X is CH or N;

P is phenyl or a 5 or 6 membered heteroaromatic ring containing one or more heteroatoms selected from N, O or S and said phenyl ring or 5 or 6 membered heteroaromatic ring may optionally be fused with a 5 or 6 membered saturated, partially saturated or unsaturated ring containing atoms independently selected from C, N, O or S;

R is CHO, fluoromethoxy, difluoromethoxy, trifluoromethoxy, C_{0-6} alkyl(SO₂)NR¹R², C_{0-6} alkyl(SO₂)NR¹R², C_{0-6} alkyl(SO₂)NR¹R², C_{0-6} alkyl(SO₂)NR¹R², C_{0-6} alkyl(SO₂)NR¹R², C_{0-6} alkylNR¹(SO₂)R², C_{0-6} alkylNR¹(SO₂)R¹R², C_{0-6} alkylNR¹(SO₂)R¹R², C_{0-6} alkyl(SO₂)C₁₋₆alkylNR¹R², C_{0-6} alkyl(SO₂)C₁₋₆alkylNR¹R², C_{0-6} alkyl(SO₂)C₁₋₆alkylNR¹R², C_{0-6} alkylNR¹R², C_{0-6} alkylNR¹R², C_{0-6} alkylNR¹R², C_{0-6} alkylNR¹R², C_{0-6} alkylOC₁₋₆alkylNR¹R², C_{0-6} alkylOC₁₋₆alkylNR¹R², C_{0-6} alkylCONR¹⁰R¹¹, C_{0-6} alkylCONR¹⁰R¹², C_{0-6} alkylNR¹¹(CO)R¹⁰, C_{0-6} alkylNR¹⁰(CO)R¹¹, C_{0-6} alkylNR¹¹(CO)R¹¹, C_{0-6} alkylNR¹⁰R¹¹, C_{0-6} alkylNR¹⁰R¹¹, C_{0-6} alkylO(CO)R¹¹, C_{0-6} AlkylO(C

 $C_{0\text{-}6}alkylC(NR^{11})N(R^{10})_2, OC_{0\text{-}6}alkylC(NR^1)NR^1R^2, C_{0\text{-}6}alkylNR^{10}(CO)OR^{11}, \\ OC_{1\text{-}6}alkylNR^1(CO)OR^2, C_{0\text{-}6}alkylNR^{11}(CO)OR^{10}, OC_{1\text{-}6}alkylCN, NR^1OR^2, C_{0\text{-}6}alkyl(CO)OR^8, \\ OC_{1\text{-}6}alkyl(CO)OR^1, NR^1(CO)NR^1R^2, NR^1(CO)(CO)R^2, NR^1(CO)(CO)NR^1R^2, OR^{12} \text{ or } SO_3R^1; \\ R^1 \text{ and } R^2 \text{ are independently selected from hydrogen, } C_{1\text{-}6}alkyl, C_{2\text{-}6}alkenyl, C_{2\text{-}6}alkynyl, \\ C_{0\text{-}6}alkylC_{3\text{-}6}cycloalkyl, C_{0\text{-}6}alkylheterocycloalkyl, C_{1\text{-}6}alkylNR^6R^7, C_{0\text{-}6}alkylaryl \text{ and } \\ C_{0\text{-}6}alkylheteroaryl, \text{ wherein any } C_{1\text{-}6}alkyl, C_{2\text{-}6}alkenyl, C_{2\text{-}6}alkynyl, C_{0\text{-}6}alkylC_{3\text{-}6}cycloalkyl, \\ C_{0\text{-}6}alkylheterocycloalkyl, C_{0\text{-}6}alkylaryl, C_{0\text{-}6}alkylheteroaryl \text{ may be substituted by one or more } A;$

R¹ and R² may together form a substituted 5 or 6 membered heterocyclic ring containing one or more heteroatoms independently selected from N, O or S, which heterocyclic ring may be optionally substituted by A;

 R^3 is independently selected from halogen, nitro, CHO, C_{0-6} alkylCN, OC_{1-6} alkylCN, C_{0-6} alkylOR 6 , OC $_{1-6}$ alkylOR 6 , fluoromethyl, difluoromethyl, trifluoromethyl, fluoromethoxy, difluoromethoxy, trifluoromethoxy, C_{0-6} alkylNR 6 R 7 , OC $_{1-6}$ alkylNR 6 R 7 , OC $_{1-6}$ alkylOC $_{1-6}$ alkylNR 6 R 7 , NR 6 OR 7 , CO $_{0-6}$ alkylCO $_{2}$ R 6 , OC $_{1-6}$ alkylCO $_{2}$ R 6 , CO $_{0-6}$ alkylCONR 6 R 7 , OC $_{1-6}$ alkylCONR 6 R 7 , OC(CO)NR 6 R 7 , OC(O)OR 6 , OC(O)R 6 , OC(O)R 6 , OC $_{1-6}$ alkylCOR 6 , OC $_{1-6}$ alkylNR 6 (CO)(CO)NR 6 R 7 , OC $_{1-6}$ alkyl(SO)NR 6 R 7 , OC $_{1-6}$ alkylNR 6 (SO)NR 6 R 7 , OC $_{1-6}$ alkylNR 6 (SO)NR 6 R 7 , OC $_{1-6}$ alkylNR 6 (SO)NR 6 R 7 , OC $_{1-6}$ alkylNR 6 (SO)NR 6 R 7 , OC $_{1-6}$ alkylNR 6 (SO)NR 7 , OC $_{1-6}$ alkylNR 6 (SO)R 7 , OC $_{1-6}$ alkylNR 7 , OC $_{1-6}$

R⁶ and R⁷ are independently selected from hydrogen, C₁-₆alkyl, C₂-₆alkenyl, C₂-₆alkynyl, C₀-₆alkylC₃-₆cycloalkyl, C₀-₆alkylaryl, C₀-₆alkylheteroaryl and C₁-₆alkylNR⁸R⁹; R⁶ and R⁷ may together form a substituted 5 or 6 membered heterocyclic ring containing one or more heteroatoms independently selected from N, O or S, which heterocyclic ring may be optionally substituted by A and wherein a CH₂ group may optionally be replaced by a CO group;

 R^8 and R^9 are independently selected from hydrogen, $C_{1\text{-}6}$ alkyl, $C_{2\text{-}6}$ alkenyl, $C_{2\text{-}6}$ alkynyl, $C_{0\text{-}6}$ alkyl $C_{3\text{-}6}$ cycloalkyl, $C_{0\text{-}6}$ alkylaryl and $C_{0\text{-}6}$ alkylheteroaryl;

R⁸ and R⁹ may together form a 5 or 6 membered heterocyclic ring containing one or more heteroatoms selected from N, O or S, which heterocyclic ring may be optionally substituted by A;

 R^{10} is hydrogen, $C_{1\text{-}6}$ alkyl, $C_{2\text{-}6}$ alkenyl, $C_{2\text{-}6}$ alkynyl, $C_{0\text{-}6}$ alkyl $C_{3\text{-}6}$ cycloalkyl, $C_{0\text{-}6}$ alkylaryl, $C_{0\text{-}6}$ alkylheteroaryl or $C_{1\text{-}6}$ alkyl NR^8R^9 ;

R¹¹ is C₁-6alkylNR⁸R⁹;

R¹⁰ and R¹¹ may together form a 5 or 6 membered heterocyclic ring containing one or more heteroatoms selected from N, O or S, which heterocyclic ring may be optionally substituted by A;

A is halogen, oxo (=O), nitro, CHO, CN, OR^6 , $C_{1\text{-}6}$ alkyl, $C_{2\text{-}6}$ alkenyl, $C_{2\text{-}6}$ alkynyl, $C_{0\text{-}6}$ alkyl $C_{3\text{-}6}$ cycloalkyl, fluoromethyl, difluoromethyl, trifluoromethyl, fluoromethoxy, difluoromethoxy, trifluoromethoxy, $C_{0\text{-}6}$ alkyl NR^6R^7 , $OC_{1\text{-}6}$ alkyl NR^6R^7 , CO_2R^8 , $CONR^6R^7$, $NR^6(CO)R^6$, $O(CO)R^6$, COR^6 , SR^6 , $(SO_2)NR^6R^7$, $(SO)NR^6R^7$, SO_3R^6 , SO_2R^6 or SOR^6 ; m is 0, 1, 2, 3 or 4;

as a free base or a salt, solvate or solvate of a salt thereof.

32. (withdrawn) A compound according to claim 31, wherein:

X is N:

P is phenyl;

R is C_{0-6} alkyl(SO_2) NR^1R^2 ;

R¹ and R² may together form a substituted 5 or 6 membered heterocyclic ring containing one or more heteroatoms independently selected from N, O or S;

m is 0:

as a free base or a salt, solvate or solvate of a salt thereof.

33. (withdrawn) A compound which is

Methyl 3-{[2,6-dimethoxy-4-(2-phenylethoxy)benzyl]amino}-6-[4-(pyrrolidin-1-ylsulfonyl)phenyl]pyrazine-2-carboxylate polystyrene;
3-{[2,6-Dimethoxy-4-(2-phenylethoxy)benzyl]amino}-6-[4-(pyrrolidin-1-ylsulfonyl)phenyl]pyrazine-2-carboxylic acid polystyrene;
as a free base or a salt, solvate or solvate of a salt thereof.

34. (Cancelled).